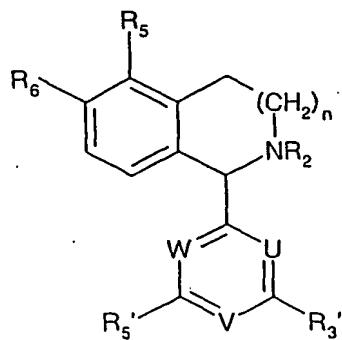


CLAIMS

1. A compound of the following general formula (I):



(I)

5 wherein

R₂ designates hydrogen, Me, Et, CHO, CN, OH, OMe, COR₉, COOR₉, CONHR₉ or CSNHR₉, whereby R₉ denotes (C₁-C₄)alkyl;

R₅ designates hydrogen, (C₁-C₄)alkyl, OH, (C₁-C₄)alkoxy, OCF₃, trifluoromethyl or halogen;

10 R₆ designates Me, (C₁-C₄)alkoxy, OCF₃, SMe or SET;

n is 1 or 2;

R₃' and R₅' each independently designate OH, Me, Et, OMe, OCF₃, trifluoromethyl or halogen;

15 U designates N or CR₂', whereby R₂' denotes hydrogen, (C₁-C₄)alkyl, (C₁-C₄)alkoxy, trifluoromethyl or halogen;

V designates N or CR₄', whereby R₄' denotes hydrogen, (C₁-C₆)alkoxy, (C₁-C₆)alkyl, OH, trifluoromethyl or halogen;

W designates N or CR₆', whereby R₆' denotes hydrogen, (C₁-C₄)alkyl, (C₁-C₄)alkoxy, trifluoromethyl or halogen;

20 and pharmaceutically acceptable salts thereof.

2. A compound according to claim 1, wherein R₂ design-

nates Me, OH, CN, CHO, COR₉ or COOR₉.

3. A compound according to claim 1, wherein R₂ designates Me, CN, CHO or COMe.

5

4. A compound according to one of claims 1 to 3, wherein R₅ designates hydrogen, Me, OMe or halogen.

10 5. A compound according to one of claims 1 to 4, wherein R₆ designates OMe or OEt.

15 6. A compound according to one of claims 1 to 3, wherein R₅ designates hydrogen or OMe, preferably hydrogen; and R₆ designates OMe.

15

7. A compound according to one of claims 1 to 6, wherein R₃' and R₅' each independently designate chloro, bromo, Me or OMe.

20 8. A compound according to claim 1 to 7, wherein R₃' and R₅' are identical; or R₃' designates chloro or bromo, and R₅' designates OMe.

25 9. A compound according to claim 7, wherein R₃' and R₅' designate both chloro or both bromo.

10. A compound according to one of claims 1 to 9, wherein U and W designate CH and V designates CR₄'.

30 11. A compound according to claim 10, wherein R₄' designates hydrogen, chloro, bromo, Me or OMe.

12. A compound according to claim 10, wherein R_3' , R_4' and R_5' designate OMe; or R_3' designates chloro and R_4' and R_5' designate OMe; or R_4' designates hydrogen and R_3' and R_5' designate both chloro or both bromo.

5 13. A compound according to claim 1, which is 1-(3,5-dichlorophenyl)-2-formyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dichlorophenyl)-2-acetyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dichlorophenyl)-2-cyano-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromo-phenyl)-2-formyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-acetyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-cyano-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-formyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-acetyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-cyano-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-formyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-acetyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-cyano-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3-chloro-4,5-dimethoxyphenyl)-2-formyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3-chloro-4,5-dimethoxyphenyl)-2-acetyl-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3-chloro-4,5-dimethoxyphenyl)-2-cyano-6-methoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dichlorophenyl)-2-formyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dichlorophenyl)-2-acetyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dichlorophenyl)-2-cyano-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-formyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-acetyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-

cyano-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-formyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-acetyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-cyano-6-ethoxy-
5 1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-formyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-acetyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-cyano-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3-chloro-4,5-dimethoxyphenyl)-2-
10 formyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3-chloro-4,5-dimethoxyphenyl)-2-acetyl-6-ethoxy-1,2,3,4-tetrahydroisoquinoline or 1-(3-chloro-4,5-dimethoxyphenyl)-2-cyano-6-ethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dichlorophenyl)-2-formyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dichlorophenyl)-2-acetyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dichlorophenyl)-2-cyano-5,6-dimethoxy-
15 1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-formyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-acetyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-cyano-5,6-dimethoxy-
20 1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dibromophenyl)-2-cyano-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-formyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-acetyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,5-dimethoxyphenyl)-2-cyano-5,6-dimethoxy-
25 1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-formyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-acetyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3,4,5-trimethoxyphenyl)-2-cyano-5,6-dimethoxy-
30 1,2,3,4-tetrahydroisoquinoline 1-(3-chloro-4,5-dimethoxyphenyl)-2-formyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline, 1-(3-chloro-4,5-dimethoxyphenyl)-2-acetyl-5,6-dimethoxy-1,2,3,4-tetrahydroisoquinoline or 1-(3-chloro-4,5-

dimethoxyphenyl)-2-cyano-5,6-dimethoxy-1,2,3,4-tetrahydroiso-quinoline, or a pharmaceutically acceptable salt thereof.

14. A compound according to anyone of claims 1 to 13, which is the (R)- or (S)-enantiomer.

5 15. A compound as defined in one of claims 1 to 14, for use as a medicament.

10 16. Use of a compound as defined in one of claims 1 to 14, for the preparation of a medicament for the prophylaxis or treatment of a disease in which down-regulation or inhibition of the expression or function of the IGF-1 receptor is beneficial.

15 17. The use according to claim 16, wherein the disease is selected from cell proliferate diseases such as cancer, atherosclerosis, restenosis, inflammatory diseases such as psoriasis, autoimmune diseases such as rheumatoid arthritis, and transplant rejection.

20 18. A method of treatment or prophylaxis of a disease in which down-regulation or inhibition of the expression or function of the IGF-1 receptor is beneficial, in a subject in need of such treatment or prophylaxis, comprising administering to said subject an amount of a compound (I) as defined in one of claims 1 to 14 in an amount which is effective in down-regulating or inhibiting the expression or function of the IGF-1 receptor.

25 19. The method of claim 18, wherein the disease is selected from cell proliferate diseases such as cancer, athero-

sclerosis, restenosis, inflammatory diseases such as psoriasis, autoimmune diseases such as rheumatoid arthritis, and transplant rejection.

20. A pharmaceutical composition comprising a compound of formula (I), or a pharmaceutically acceptable salt thereof as defined in any one of claims 1 to 14, and a pharmaceutically acceptable adjuvant, diluent or carrier.

21. Articles containing a compound of the formula (I) or a pharmaceutically acceptable salt thereof as defined in any one of claims 1 to 14, and a chemotherapeutic agent, as a combination for the simultaneous, separate or successive administration in the therapy of a disease in which down-regulation or inhibition of the expression or function of the IGF-1 receptor is beneficial.

22. Use of a compound of the formula (I) or a pharmaceutically acceptable salt thereof as defined in any of claims 1 to 14, as a pharmacological tool in the development and standardization of in vitro and in vivo test systems for the evaluation of the effects of inhibitors of cell cycle activity in laboratory animals.